

ORDER FOR SUPPLIES OR SERVICES

PAGE OF PAGES

1

30

IMPORTANT: Mark all packages and papers with contract and/or order numbers.

1. DATE OF ORDER 06/08/2021		2. CONTRACT NO. (If any) 68HE0H18D0008		6. SHIP TO: a. NAME OF CONSIGNEE Office of Research and Development	
3. ORDER NO. 68HERC21F0259		4. REQUISITION/REFERENCE NO. PR-ORD-20-02465		b. STREET ADDRESS US Environmental Protection Agency 109 T.W. Alexander Drive	
5. ISSUING OFFICE (Address correspondence to) CAD US Environmental Protection Agency 26 West Martin Luther King Drive Mail Code: W136 Cincinnati OH 45268-0001		c. CITY Research Triangle Park		d. STATE NC	e. ZIP CODE 27711
7. TO: Linda Eubanks		f. SHIP VIA			
a. NAME OF CONTRACTOR Research Triangle Institute		8. TYPE OF ORDER <input type="checkbox"/> a. PURCHASE REFERENCE YOUR: vendor proposal dated April 12, 2021. Please furnish the following on the terms and conditions specified on both sides of this order and on the attached sheet, if any, including delivery as indicated.			
b. COMPANY NAME		<input checked="" type="checkbox"/> b. DELIVERY Except for billing instructions on the reverse, this delivery order is subject to instructions contained on this side only of this form and is issued subject to the terms and conditions of the above-numbered contract.			
c. STREET ADDRESS PO BOX 12194					
d. CITY RESEARCH TRIANGLE PARK	e. STATE NC	f. ZIP CODE 277092194			
9. ACCOUNTING AND APPROPRIATION DATA See Schedule		10. REQUISITIONING OFFICE Office of Research and Development			

11. BUSINESS CLASSIFICATION (Check appropriate box(es)) <input type="checkbox"/> a. SMALL <input checked="" type="checkbox"/> b. OTHER THAN SMALL <input type="checkbox"/> c. DISADVANTAGED <input type="checkbox"/> d. WOMEN-OWNED <input type="checkbox"/> e. HUBZone <input type="checkbox"/> f. SERVICE-DISABLED VETERAN-OWNED <input type="checkbox"/> g. WOMEN-OWNED SMALL BUSINESS (WOSB) ELIGIBLE UNDER THE WOSB PROGRAM <input type="checkbox"/> h. EDWOSB				12. F.O.B. POINT	
13. PLACE OF a. INSPECTION Destination		b. ACCEPTANCE Destination		14. GOVERNMENT B/L NO.	
				15. DELIVER TO F.O.B. POINT ON OR BEFORE (Date) 06/08/2024	
				16. DISCOUNT TERMS	

17. SCHEDULE (See reverse for Rejections)

ITEM NO. (a)	SUPPLIES OR SERVICES (b)	QUANTITY ORDERED (c)	UNIT (d)	UNIT PRICE (e)	AMOUNT (f)	QUANTITY ACCEPTED (g)
	DUNS Number: 004868105+7709 Endocrine Disruptor Screening Program Technical and Support Services ID/IQ Task Order: Laboratory Support for Repeated Dose Oral Toxicity Studies of Poly- and Continued ...					
18. SHIPPING POINT		19. GROSS SHIPPING WEIGHT		20. INVOICE NO.		17(h) TOTAL (Cont. pages)
21. MAIL INVOICE TO: a. NAME RTP Finance Center				\$4,527,202.00		
b. STREET ADDRESS (or P.O. Box) US Environmental Protection Agency RTP-Finance Center (AA216-01) 109 TW Alexander Drive www2.epa.gov/financial/contracts		c. CITY Durham		d. STATE NC	e. ZIP CODE 27711	17(i) GRAND TOTAL

22. UNITED STATES OF AMERICA BY (Signature)	06/08/2021	23. NAME (Typed) Sean Gifford TITLE: CONTRACTING/ORDERING OFFICER
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ORDER FOR SUPPLIES OR SERVICES
SCHEDULE - CONTINUATION

PAGE NO
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IMPORTANT: Mark all packages and papers with contract and/or order numbers.

DATE OF ORDER 06/08/2021	CONTRACT NO. 68HE0H18D0008	ORDER NO. 68HERC21F0259
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ITEM NO. (a)	SUPPLIES/SERVICES (b)	QUANTITY ORDERED (c)	UNIT (d)	UNIT PRICE (e)	AMOUNT (f)	QUANTITY ACCEPTED (g)
0001	<p>Perfluoroalkyl Substances in Rats TOCOR: Patrice Borsz Max Expire Date: 06/08/2024 Invoice Approver: Patrice Borsz Alt Invoice App: Jenna Tomayko Admin Office: CAD US Environmental Protection Agency 26 West Martin Luther King Drive Mail Code: W136 Cincinnati OH 45268-0001 Period of Performance: 06/09/2021 to 06/08/2024</p> <p>Base Period (Tasks 1 and 3) - In accordance with Attachment 1, Performance Work Statement.</p> <p>This Contract Line Item (CLIN) is Time & Materials (T&M), with an associated not-to-exceed ceiling of \$593,951.00.</p> <p>The associated period of performance for this CLIN is from June 9, 2021 to June 8, 2024.</p> <p>Accounting Info: 20-21-C3-26B2000-000FK9-2532-HQ00BG00- 26A5C-2026B2E055-008 BFY: 20 EFY: 21 Fund: C3 Budget Org: 26B2000 Program (PRC): 000FK9 Budget (BOC): 2532 Job #: HQ00BG00 Cost: 26A5C DCN - Line ID: 2026B2E055-008 Funding Flag: Complete Funded: \$593,951.00</p>				593,951.00	
0002	<p>Base Period (Tasks 2, 4, and 5) in accordance with Attachment 1, Performance Work Statement.</p> <p>This is a firm-fixed priced contract line item number (CLIN).</p> <p>This CLIN has an associated period of performance from June 9, 2021 to June 8, 2024.</p> <p>Continued ...</p>				3,215,702.00	

TOTAL CARRIED FORWARD TO 1ST PAGE (ITEM 17(H))

\$3,809,653.00

ORDER FOR SUPPLIES OR SERVICES
SCHEDULE - CONTINUATION

PAGE NO
3

IMPORTANT: Mark all packages and papers with contract and/or order numbers.

DATE OF ORDER 06/08/2021	CONTRACT NO. 68HE0H18D0008	ORDER NO. 68HERC21F0259
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ITEM NO. (a)	SUPPLIES/SERVICES (b)	QUANTITY ORDERED (c)	UNIT (d)	UNIT PRICE (e)	AMOUNT (f)	QUANTITY ACCEPTED (g)
0003	<p>Accounting Info: 20-21-C3-26B2000-000FK9-2532-HQ00BG00-26A5C-2026B2E055-008 BFY: 20 EFY: 21 Fund: C3 Budget Org: 26B2000 Program (PRC): 000FK9 Budget (BOC): 2532 Job #: HQ00BG00 Cost: 26A5C DCN - Line ID: 2026B2E055-008 Funding Flag: Complete Funded: \$1,839,881.00</p> <p>Accounting Info: 21-22-C3-26B2000-000FK9-2532-HQ00BG00-26A5C-2026B2E055-007 BFY: 21 EFY: 22 Fund: C3 Budget Org: 26B2000 Program (PRC): 000FK9 Budget (BOC): 2532 Job #: HQ00BG00 Cost: 26A5C DCN - Line ID: 2026B2E055-007 Funding Flag: Complete Funded: \$1,375,821.00</p> <p>Base Period (Optional Task 6) in accordance with Attachment 1, Performance Work Statement.</p> <p>This is a firm-fixed priced contract line item number (CLIN).</p> <p>This CLIN has an associated period of performance from June 9, 2021 to June 8, 2024.</p> <p>This optional CLIN is exercised upon award of the task order.</p> <p>Accounting Info: 20-21-C-26B2000-000FK9XR6-2532-26A5C-2026B2E055-005 BFY: 20 EFY: 21 Fund: C Budget Org: 26B2000 Program (PRC): 000FK9XR6 Budget (BOC): 2532 Cost: 26A5C DCN - Line ID: 2026B2E055-005 Funding Flag: Complete Funded: \$2,981.00</p> <p>Accounting Info: 20-21-C-26B2000-000FK9XR5-2532-26A5C-2026B2E055-002 BFY: 20 EFY: 21 Fund: C Budget Org: 26B2000 Program (PRC): 000FK9XR5 Budget (BOC): 2532 Cost: Continued ...</p>				717,549.00	

TOTAL CARRIED FORWARD TO 1ST PAGE (ITEM 17(H))

\$717,549.00

ORDER FOR SUPPLIES OR SERVICES
SCHEDULE - CONTINUATION

PAGE NO
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IMPORTANT: Mark all packages and papers with contract and/or order numbers.

DATE OF ORDER 06/08/2021	CONTRACT NO. 68HE0H18D0008	ORDER NO. 68HERC21F0259
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ITEM NO. (a)	SUPPLIES/SERVICES (b)	QUANTITY ORDERED (c)	UNIT (d)	UNIT PRICE (e)	AMOUNT (f)	QUANTITY ACCEPTED (g)
	26A5C DCN - Line ID: 2026B2E055-002 Funding Flag: Complete Funded: \$6,680.00 Accounting Info: 20-21-C-26B2000-000FK8XPW-2532-26A5C-2 026B2E055-003 BFY: 20 EFY: 21 Fund: C Budget Org: 26B2000 Program (PRC): 000FK8XPW Budget (BOC): 2532 Cost: 26A5C DCN - Line ID: 2026B2E055-003 Funding Flag: Complete Funded: \$133.00 Accounting Info: 20-21-C-26B2000-000FK7XR4-2532-26A5C-2 026B2E055-004 BFY: 20 EFY: 21 Fund: C Budget Org: 26B2000 Program (PRC): 000FK7XR4 Budget (BOC): 2532 Cost: 26A5C DCN - Line ID: 2026B2E055-004 Funding Flag: Complete Funded: \$324.00 Accounting Info: 21-22-C-26B2000-000FK9XR6-2532-26A5C-2 026B2E055-006 BFY: 21 EFY: 22 Fund: C Budget Org: 26B2000 Program (PRC): 000FK9XR6 Budget (BOC): 2532 Cost: 26A5C DCN - Line ID: 2026B2E055-006 Funding Flag: Complete Funded: \$4,541.00 Accounting Info: 21-22-C3-26B2000-000FK9-2532-HQ00BG00- 26A5C-2026B2E055-007 BFY: 21 EFY: 22 Fund: C3 Budget Org: 26B2000 Program (PRC): 000FK9 Budget (BOC): 2532 Job #: HQ00BG00 Cost: 26A5C DCN - Line ID: 2026B2E055-007 Funding Flag: Complete Funded: \$702,890.00					

TOTAL CARRIED FORWARD TO 1ST PAGE (ITEM 17(H))

\$0.00



ENVIRONMENTAL PROTECTION AGENCY

EPA Office of Research and Development: Laboratory Support for Repeated Dose Oral Toxicity Studies of Poly- and Perfluoroalkyl Substances in Rats

Task Order 68HERC21F0259

Version 1, Dated 06/07/2021

1 PERFORMANCE WORK STATEMENT (PWS)

1.1 PURPOSE

The purpose of this task order (TO), *EPA Office of Research and Development: Laboratory Support for Repeated Dose Oral Toxicity Studies of Perfluoroalkyl Substances (PFAS) in Rats*, is to obtain laboratory support for the Office of Science Coordination and Policy (OSCP), Office of Chemical Safety & Pollution Prevention (OCSPP) and Office of Research and Development in four (4) general areas:

1. Determine purity, and solubility and stability of selected PFAS chemicals in dosing solution;
2. Tolerability and 14-day oral dose range finding study of selected PFAS chemicals;
3. 90-day oral subchronic toxicity study of selected PFAS chemicals; and
4. 28-day Immunotoxicity Study for Perfluoroalkyl Substances.

The work under this task order falls within scope of the contract level statement of work under Section 3.1 Laboratory Testing using EPA or OECD Test Guidelines (Task 1), specifically, 3.1.2 Tier 1 and Tier 2 in vivo mammalian guideline tests (see in vivo assays in Table 1).

Animal Care

1. Animal Welfare Oversight: Vertebrate animal studies shall be conducted by a PHS Assured and AAALAC, Int. Accredited institution using an Institutional Care and Use Committee (IACUC) approved protocol. The ownership and welfare of the animals shall be the responsibility of this Assured and Accredited institution. Assurance number and Accreditation Unit shall be provided to the EPA.
2. Animal Husbandry will follow the requirements of the Guide for the Care and Use of Laboratory Animals. Animals shall be socially housed 2 per cage. The temperature in the experimental animal room shall be 22°C (\pm 3°C) with a relative humidity of 50 \pm 20 percent. Lighting shall be artificial with a sequence of 12 hours light, 12 hours dark. Diet and water shall be provided ad libitum.

1.2 BACKGROUND

The Office of Research and Development (ORD) provides scientific research data to inform EPA's regulatory decisions and support its partners and stakeholders including the EPA Regional offices, State Environmental and Public Health programs and tribal organizations.

Per/polyfluoroalkyl substances (PFAS) are used in industrial processes and found in commercial products. Through their manufacture, use and disposal, these compounds have become anthropogenic contaminants in air, soil and water. These chemicals tend to be bioaccumulative and persistent in the environment. Several of these compounds, of which there may be thousands, have been detected in human blood. Some PFAS compounds are associated with chronic effects such as immunotoxicity and potentially cancer. ORD's Center for Computational Toxicology and Exposure (CCTE) is currently conducting high-throughput in vitro toxicity assays with many of these PFAS chemicals. However, for many of the PFAS compounds, the available data on the potential in vivo toxic effects from acute and chronic exposure are scant. In order to provide an in vivo anchor for the in vitro toxicity assay results and read-across approaches being developed, CCTE is requesting proposals to conduct repeated oral dose toxicity assays on a limited number of chemicals. The oral route was chosen as this is anticipated to be the major route of exposure for the general population, either through contaminated drinking water or food. The overall objective of this task order is to evaluate the 90-day oral toxicity of selected PFAS compounds in the rat. To achieve this objective, the contractor shall: 1) determine the purity of EPA-selected PFAS compounds and their solubility and stability in vehicle; 2) conduct tolerability and 14-day

oral dose range finding studies; 3) conduct 90-day oral toxicity studies; and 4) report the findings to the EPA. Studies may be conducted on up to six compounds. Preferably, the studies shall be conducted in parallel. For each chemical, an option to perform an additional 28-day oral immunotoxicity study may be requested.

1.3 TASK 1: TASK ORDER MANAGEMENT AND REPORTING REQUIREMENTS

Task 1 will be Time & Materials.

- 1.3.1 The Contractor shall schedule a kick-off meeting/conference call with the TOCOR within 10 business days following the TO award. The TOCOR, Alternate (Alt) TOCOR, contract-level COR, and the EPA Contracting Officer (CO) must be invited to the kick-off meeting. Additional participants may be included.
- 1.3.2 The Contractor shall manage all aspects of the task order including, but not limited to, the technical, quality assurance, schedule, cost, and communication requirements.
- 1.3.3 General status meetings shall be conducted on an as needed basis and may be initiated by the TOCOR or contractor. Meetings may be required at the end of each deliverable to discuss the deliverable, progress of the overall project, problems encountered and next steps. Meetings may be held in-person or via teleconference technology such as Skype, MS Teams, Adobe Connect, etc. It is expected that much of the day-to-day communication for this task order will be conducted via email.
- 1.3.4 The Contractor shall update the TOCOR via telephone and, in writing, via e-mail, of any issues on an ongoing basis.
- 1.3.5 The Contractor shall immediately inform the TOCOR when any hours or costs for time and material contractual tasks (Task 3) that has exceeded or is expected to exceed the contractor estimate by >10%. This is for the procurement of chemicals and analytical chemistry (Task 3). Tasks 4, 5 and 6 are firm-fixed price.
- 1.3.6 The Contractor shall immediately inform the TOCOR of any problems that may impact the production, budget, and/or delivery of deliverables.
- 1.3.7 The Contractor shall notify the TOCOR when 75% of the Government approved hours or approved LH (labor hour) costs have been incurred (including unbilled hours and costs).
- 1.3.8 The Contractor shall provide a monthly progress report of the combined monthly technical and financial progress report) stating the progress made, including the percentage of the project completed, a description of the work accomplished to support the cost, the estimated percentage of task completed (including deliverables) during the reporting period. The Executive Summary shall summarize the planned and actual work for the month, financial status, work planned for the next month, and significant issues, risks, or concerns. The monthly report shall also provide cost and technical progress data for each time and materials task and projected costs for the upcoming reporting period.

1.3.9 For the technical progress report also include the following specific information:

- Narrative detail review of accomplishments during the reporting period and/or significant events, as well as an assessment of work being completed on schedule and budget.
- Status of all ongoing activities in accordance with the technical proposal and technical directives.
- List of deliverables with delivery dates (planned versus actual).
- Anticipated activities and deliverables for the next reporting period.
- Specific discussions shall include difficulties encountered and remedial action taken during the reporting period, and anticipated activity with a schedule of deliverables for the subsequent reporting period.
- List of current contractors / staffing roster and any changes that may impact deliverables in advance of the reporting period (e.g., change in personnel and vacations).
- Monthly Contractor performance information (performance metrics)

1.3.10 For the Monthly Progress Report with regard to Task 3, include the following information:

- Identification of cost issues or concerns.
- For the current period, display the amount claimed.
- For the cumulative period display the total amount claimed; amount paid; amount suspended or disallowed; and remaining amount.
- Labor hours
 - A list of employees, their labor categories, and the number of hours worked for the reporting period.
 - For the current reporting period display the expended direct labor hours (by EPA contract labor category), and the total loaded direct labor hours.
 - For the cumulative reporting period and the cumulative contract period display: The negotiated and expended direct labor hours (by EPA labor hour category) and the loaded direct labor rate.
 - Display the estimated direct labor hours and costs to be expended during the next reporting period.

- Display the estimates of remaining direct labor hours and costs required to complete the task order.
 - Unbilled allowable costs. Display the total costs incurred but unbilled for the current reporting period and cumulative for the task order.
 - Average total cost labor hour. For the current contract period, compare the actual total cost per hour to date with the average total cost per hour of the approved technical proposal for the task order.
 - The monthly report does not change the notification requirements of the “Limitation of Cost” or “Limitation of Funds” clauses requiring separate written notice to the Contracting Officer.
- 1.3.11 The Contractor shall maintain a cumulative record of all communications between the contractor and EPA (all media including e-mail and telephone calls) and provide it to the TOCOR within one month after the TO has ended.
- 1.3.12 The contractor shall provide all deliverables in an electronic format specified by the EPA TOCOR (e.g., Word, Excel, Access, HTML) via electronic mail. The Contractor shall format any deliverables intended for posting on an EPA public website to comply with Section 508.
- 1.3.13 Unless otherwise specified by the TOCOR, the Contractor shall provide a secure method for internet transfer of large files.
- 1.3.14 All deliverables for this task order are the property of EPA.
- 1.3.15 Contractor personnel shall identify themselves as contractor employees and shall not present themselves as EPA employees. Furthermore, they shall not represent view of the U.S. Government, EPA, or its employees. In addition, the contractor shall not engage in inherently governmental activities, including, but not limited to actual determination of EPA policy and preparation of documents on EPA letterhead other than routine correspondences.

1.4 TASK 2: QUALITY ASSURANCE AND QUALITY ASSURANCE PROJECT PLAN (QAPP)

The Contractor shall implement a quality system that meets ANSI standard E4-2014.

Task 2 will be Firm-Fixed Price.

For planning purposes, assume that a Quality Assurance Project Plan (QAPP) will be required for Tasks 3, 4 and 5. The contractor shall create a Quality Assurance Project Plan (QAPP) that documents the planning, implementation, and assessment procedures for quality assurance and quality control activities. The QAPP integrates all the technical and quality aspects of the project to provide a blueprint for obtaining the type and quality of data and information needed for a specific decision or use. The QAPP shall be prepared in accordance with the specifications identified by EPA (found at <https://www.epa.gov/quality/epa-qar-5-epa-requirements-quality-assurance-project-plans>).

- Within 10 business days after Task Order Award, the contractor shall prepare and submit for EPA review a draft Quality Assurance Project Plan (QAPP) for Tasks 3, 4 and 5. The QAPP shall be submitted to the TO COR.
- EPA will review the contractor's draft QAPP and provide the Contractor with written approval or written comments.
- If needed, the Contractor shall submit a revised QAPP within 5 business days of receipt of the written comments on the draft QAPP, unless otherwise instructed by the EPA TO COR.
- Under no circumstances shall work that involves the generation, collection, evaluation, analysis, or use of environmental data be performed by the contractor until the contractor receives written notification from the EPA TO COR that EPA has approved the contractor's QAPP.
- All QA documentation, including the QAPP, prepared under this Task Order, shall be considered non-proprietary, and shall be made available to the public upon request.
- The contractor also shall provide EPA with monthly reports of QA-related activities performed during implementation of this Task Order. These monthly QA reports shall identify QA activities performed to support implementation of this task order, problems encountered, deviations from the QAPP, and corrective actions taken.

1.5 TASK 3: CHEMICAL PROCUREMENT AND ANALYTICAL CHEMISTRY

Purpose: The purpose of Task 3 is for the contractor to purchase selected PFAS chemicals and to develop analytical chemistry methods to determine the purity, and solubility and stability of the PFAS chemicals in dosing vehicle.

Task 3 will be Time and Materials

Task 3.1: Chemical Procurement

The following chemicals shall be used in the study:

Table 1

DTXSID	Name	CAS RN
DTXSID00194615	1H,1H,9H-Perfluorononyl acrylate	4180-26-1
DTXSID1047578	1H,1H,2H,2H-Perfluorohexyl iodide	2043-55-2
DTXSID9059834	Perfluamine	338-83-0
DTXSID00188993	Methyl heptafluoropropylketone	355-17-9
DTXSID 3031860	Nonadecafluorodecanoic acid	335-76-2
DTXSID6067331	1H,1H,2H,2H-Perfluorooctanesulfonic acid	27619-97-2

A. Chemical Procurement

The contractor shall purchase the chemical(s) selected by the EPA. Preferably, a single lot shall be used throughout the study.

B. Deliverables

Provide the EPA information regarding test chemical procured. Include purity (manufacturer), lot number, and other information provided by the manufacturer. Information provided electronically in Word format. Report shall be submitted within two weeks of acquisition of chemical.

Task 3.2: Chemical Purity

A. Chemical Purity

The contractor shall develop analytical methods (LC-MS/MS or GC-MS/MS, based on EPA recommendations) to determine the purity of the purchased chemicals. If multiple lots of the chemical are purchased, purity checks on each lot are required. The contractor must generate Standard Operating Procedures (SOPs) for the analytical chemistry methods. Purity of test chemicals shall be >95%, unless specifically approved by EPA. The levels of the potential contaminants perfluorooctanoic acid (PFOA) and perfluorooctane sulfonic acid (PFOS) shall be reported.

Task 3.3: Chemical Solubility and Stability

A. Solubility and stability of chemical in vehicle

The contractor shall determine the solubility and stability of the test compounds in appropriate vehicle using chromatographic/mass spectrometry methods. A standard set of dosing vehicles shall be evaluated for chemical solubility and stability. The vehicles may include deionized water, 1:1:8 Kolliphor:ethanol:deionized water, and deionized water with 20% Tween® 80. The solubility and stability of the chemicals in other vehicles may be assessed as needed. The stability of the chemical shall be evaluated in the dosing solution for 7 days. The dosing vehicle used in the study shall be approved by the US EPA. In some cases, chemical may be used in suspension. Deviation of concentration of test chemical in dosing solutions greater than 10% from target concentration are considered out of tolerance.

B. Deliverables

Within two (2) months of the approval of the QAPP a draft of all SOPs shall be provided to the EPA for approval. Each SOP shall have clearly labeled sections similar to: 1.0 Purpose; 2.0 Scope; 3.0 Responsibilities; 4.0 Procedures; 5.0 References. The draft SOPs shall be delivered to the EPA electronically in MS Word format and all data shall be provided in Excel spreadsheets. The EPA reserves the right to review or modify the SOPs prior to approval. SOPs must be approved before task order work can move forward into Task 4.

Within 8 weeks of the approval of the Chemistry SOP's, a draft analytical chemistry report shall be provided to the EPA for each test chemical. Each analytical chemistry report shall include the detailed analytical method, the Limit of Quantitation (LOQ) and Level of Detection (LOD), the purity of the test chemical, and the solubility and stability of the test chemical in the dosing vehicle. This analytical chemistry report shall be in a Word format with all data provided in an Excel format and must be approved by EPA before any in vivo testing (Task 4.1) shall occur. In the event that a chemical does not pass purity or stability checks, the EPA shall provide an alternative chemical.

1.6 TASK 4: ORAL TOXICITY STUDY – DOSE RANGE FINDING

Purpose: The purpose of Task 4 is for the contractor to conduct a phased non-GLP an acute oral tolerability study and a 14-day repeated dose oral toxicity study to determine doses that shall be used in the 90-day study.

Studies may be conducted on up to 6 compounds.

Task 4 will be Firm-Fixed Price

Task 4.1: Tolerability Study

A. Tolerability Study

The contractor shall conduct a non-GLP tolerability study for each chemical to be evaluated in the subchronic study. The purpose of the tolerability study is to guide dose selection for the 14-day dose range finding study. Up to six (6) tolerability studies may be conducted depending on funding. The study design is as follows:

- 1) Species and Sex: Testing shall be performed in male Sprague-Dawley IGS rats from Charles Rivers Laboratory.
- 2) Age: Testing shall be performed with clinically healthy animals following a 7 – 10-day acclimation period. Dosing shall begin with animals no later than 8 – 9 weeks of age.
- 3) Weight: at the commencement of the study the weight variation of animals used shall be within 20 percent of the mean weight for each sex.
- 4) Numbers: A total of 3 animals shall be used per dose level.
- 5) Animal Husbandry: Studies shall be conducted in a facility accredited by the Association for the Assessment and Accreditation for Laboratory Animal Laboratory Care (AAALAC). Animals shall be group housed 2 per cage. The temperature in the experimental animal room shall be 22°C (\pm 3°C) with a relative humidity of 50 \pm 20 percent. Lighting shall be artificial with a sequence of 12 hours light, 12 hours dark. Diet and water shall be provided ad libitum.
- 6) Control and Test Substance: The vehicle and test substances will be defined in Task 3.
- 7) Dose Levels: A total of 4 dose levels shall be used. Dose levels will be provided by EPA. A concurrent vehicle control group is required.
- 8) Administration of Test Substance: The test substance shall be administered as a single dose, oral gavage. A dose volume of 5 ml/kg is preferred but 10 mL/kg is acceptable if required for solubility.
- 9) Observation Period and Observations: The animals shall be observed for 72 hours following dosing. Observations shall be made at least twice each day for morbidity and mortality. General clinical observations shall be made at least twice a day, preferably at the same time each day. Effort shall be made to ensure that variations in the observation conditions are minimal. Observations shall be detailed and carefully recorded, preferably using scoring

systems, explicitly defined by the testing laboratory. Signs noted shall include, but not be limited to, changes in skin, fur, eyes, mucous membranes, occurrence of secretions and excretions and autonomic activity (e.g., lacrimation, piloerection, pupil size, unusual respiratory pattern). Changes in gait, posture and response to handling as well as the presence of clonic or tonic movements, stereotypies (e.g., excessive grooming, repetitive circling) or bizarre behavior (e.g., self-mutilation, walking backwards) shall be recorded. Measurement of food and water consumption shall be recorded over the 72-hour period. Individual weights of animals shall be determined shortly before the test substance is administered and at death or scheduled sacrifice. Moribund animals shall be removed and sacrificed when noticed and the time of death shall be recorded as precisely as possible. At termination, all survivors in the treatment and control groups shall be sacrificed.

10) Data and Reporting: Data shall be summarized in tabular form, showing for each test group the number of animals at the start of the test, the number of animals showing lesions, the types of lesions and the percentage of animals displaying each type of lesion. In addition to reporting requirements, the following specific information shall be reported:

- a. Test substance characterization shall include:
 - i. Chemical identification.
 - ii. Lot or batch number.
 - iii. Physical properties.
 - iv. Purity/impurities.
 - v. Identification and composition of any vehicle used.
- b. Test system shall contain data on:
 - i. Species and strain of animals used.
 - ii. Age including body weight data and sex.
 - iii. Test environment including cage conditions, ambient temperature, humidity, and light/dark periods.
 - iv. Identification of animal diet.
 - v. Acclimation period.
- c. Test procedure shall include the following data:
 - i. Method of randomization used.
 - ii. Full description of experimental design and procedure.

- iii. Dose regimen including levels, methods, and volume.
- d. Test results shall include:
 - i. Group animal data. Tabulation of toxic response data by sex and exposure level for:
 - 1. Number of animals exposed.
 - 2. Number of animals showing signs of toxicity.
 - 3. Number of animals dying.
 - ii. Individual animal data. Data shall be presented as summary (group mean) as well as for individual animals.
 - 1. Date of death during the study or whether animals survived to termination.
 - 2. Date of observation of each abnormal sign and its subsequent course.
 - 3. Body weight data.
 - 4. Feed and water (if collected) consumption data.
 - 5. Dose (mg/kg/day).
 - 6. Statistical treatment of results, where appropriate.

B. Deliverables

Within two (2) weeks of completion of the tolerability study (Task 4.1), the contractor shall provide a report of the study results electronically to the EPA in MS Word format and all data in Excel spreadsheets.

Task 4.2: Dose Range Finding Study

A. 14-Day Repeated Oral Dose Range Finding Study

The contractor shall conduct a non-GLP, 14-day repeated oral dose range finding study for each chemical evaluated in a subchronic toxicity study. The purpose of the dose range finding study is to guide dose selection for the subchronic toxicity study. Up to six (6) 14-day dose range studies may be conducted depending on funding. The study design is as follows:

- 1) Species and Sex: Testing shall be performed in male and female Sprague-Dawley IGS rats from Charles Rivers Laboratory.
- 2) Age: Testing shall be performed with clinically healthy animals following a 7 – 10-day acclimation period. Dosing shall begin with animals no later than 8 – 9 weeks of age.

- 3) Weight: at the commencement of the study the weight variation of animals used shall be within 20 percent of the mean weight for each sex.
- 4) Numbers: A total of 5 animals per sex per dose level shall be used.
- 5) Animal Husbandry: Studies shall be conducted in a facility accredited by the Association for the Assessment and Accreditation of Laboratory Animal Care (AAALAC). Animals shall be group housed 2 per cage. The temperature in the experimental animal room shall be 22°C (\pm 3°C) with a relative humidity of 50 \pm 20 percent. Lighting shall be artificial with a sequence of 12 hours light, 12 hours dark. Diet and water shall be provided ad libitum.
- 6) Control and Test Substance: The vehicle and test substances shall be defined in Task 3.
- 7) Dose Levels: A total of 3 dose levels shall be used. Dose levels shall be provided by EPA based on the tolerability study. The high dose shall demonstrate adverse, but survivable effects, preferably in the high-dose group only. A concurrent vehicle control group is required.
- 8) Administration of Test Substance: The test substance shall be administered via oral gavage every day for 14 consecutive days. A dose volume of 5 ml/kg is preferred but 10 mL/kg is acceptable if required for solubility. The dose shall be given at approximately the same time each day and adjusted based on daily body weights to maintain a consistent dose level.
- 9) Observation Period and Observations: The animals shall be observed for 14 days. Observations shall be made at least twice each day for morbidity and mortality. Appropriate actions shall be taken to minimize loss of animals to the study (e.g., necropsy or refrigeration of those animals found dead and isolation or sacrifice of weak or moribund animals). General clinical observations shall be made at least once per day, preferably at the same time each day. Effort shall be made to ensure that variations in the observation conditions are minimal. Observations shall be detailed and carefully recorded, preferably using scoring systems, explicitly defined by the testing laboratory. Signs noted shall include, but not be limited to, changes in skin, fur, eyes, mucous membranes, occurrence of secretions and excretions and autonomic activity (e.g., lacrimation, piloerection, pupil size, unusual respiratory pattern). Changes in gait, posture and response to handling as well as the presence of clonic or tonic movements, stereotypies (e.g., excessive grooming, repetitive circling) or bizarre behavior (e.g., self-mutilation, walking backwards) shall be recorded. Measurement of food consumption shall be recorded weekly. Individual weights of animals shall be determined each day before the test substance is administered and at death. Moribund animals shall be removed and sacrificed when noticed and the time of death shall be recorded as precisely as possible.
- 10) Sacrifice: At termination, all survivors in the treatment and control groups shall be weighed and then sacrificed. The order of sacrifice among animals shall be randomized.
- 11) Clinical Pathology: Hematology and clinical chemistry examinations shall be made on all animals, including controls, of each sex in each group. The hematology and clinical

chemistry parameters shall be examined at terminal sacrifice at the end of the study. Overnight fasting of the animals prior to blood sampling is recommended.

- a. Hematology. Measured endpoints shall be red blood cell count, hemoglobin concentration, hematocrit, mean corpuscular volume, mean corpuscular hemoglobin, and mean corpuscular hemoglobin concentration, white blood cell count, differential leukocyte count, platelet count, and a measure of clotting potential, such as prothrombin time or activated partial thromboplastin time.
 - b. Clinical chemistry. Measured endpoints shall be potassium, sodium, glucose, total cholesterol, urea nitrogen, creatinine, total protein and albumin. Hepatic enzymes alanine aminotransferase, aspartate aminotransferase, and alkaline phosphatase shall also be measured.
- 12) Plasma collection and chemical analysis: On the first day of dosing, blood shall be collected (using K3 EDTA as anticoagulant) from the retro-orbital plexus or either tail or jugular venipuncture under 70% CO₂:30% O₂ anesthesia. Plasma shall be isolated, stored at -80° C (2 x 50 µL aliquots of each). The first blood collection shall be done at approximately 2 hours post-administration. The time of collection shall be recorded. At the time of sacrifice, blood shall be collected (using K3 EDTA as anticoagulant) by cardiac puncture under 70% CO₂:30% O₂ anesthesia. Plasma shall be isolated, stored at -80° C (6 x 100 µL aliquots of each). The time of blood collection and necropsy shall be recorded for all animals. Plasma samples analyzed by contractor for parent compound and metabolites by a suitable chromatographic/mass spectrometry method. The government's primary objective of this analysis is the quantitation of parent compound in plasma. Based on the method for the parent, we would like to identify, but not quantify, any metabolites that are present.
- 13) Ophthalmological Examination: Ophthalmological examinations using an ophthalmoscope or an equivalent device shall be made on all animals prior to the administration of the test substance and on all high dose and control groups at termination. If changes in the eyes are detected, all animals in the other dose groups shall be examined.
- 14) Gross Necropsy: All animals shall be subjected to a full gross necropsy which includes examination of the external surface of the body, all orifices, and the cranial, thoracic and abdominal cavities and their contents.
- a. The liver, kidneys, lungs, adrenals, testes, epididymides, ovaries, uterus, thymus, spleen, brain, and heart shall be trimmed and weighed wet, as soon as possible after dissection.
 - b. The following organs and tissues, or representative samples thereof, shall be preserved in a suitable medium for histopathological examination:
 - i. Digestive system—salivary glands, esophagus, stomach, duodenum, jejunum, ileum, cecum, colon, rectum, liver, pancreas, gallbladder (when present).
 - ii. Nervous system—brain (including sections of medulla/pons, cerebellum and cerebrum), pituitary, peripheral nerve (sciatic or tibial, preferably in close

proximity to the muscle), spinal cord (three levels: cervical, mid-thoracic and lumbar), eyes (retina, optic nerve).

- iii. Glandular system—adrenals, parathyroid, thyroid.
- iv. Respiratory system—trachea, lungs, pharynx, larynx, nose.
- v. Cardiovascular/hemopoietic system—aorta, heart, bone marrow (and/or fresh aspirate), lymph nodes (preferably one lymph node covering the route of administration and another one distant from the route of administration to cover systemic effects), spleen, thymus.
- vi. Urogenital system—kidneys, urinary bladder, prostate, testes, epididymides, seminal vesicle(s), uterus, ovaries, female mammary gland.
- vii. Others—all gross lesions and masses, skin.

15) Histopathology: The following histopathology shall be performed:

- a. Full histopathology on the organs and tissues listed under 14(b) of all animals in the control and high dose groups. If suspected treatment-related changes in an organ or tissue are detected, all animals in the other dose groups shall be examined. If excessive early deaths or other problems occur in the high dose group compromising the significance of the data, the next dose level shall be examined for complete histopathology.
- b. Full histopathology shall be performed on the organs and tissues listed under 14(b) in all animals that died or were killed during the study.
- c. Full histopathology shall be performed on all gross lesions in all animals.
- d. An attempt shall be made to correlate gross observations with microscopic findings.
- e. Tissues and organs designated for microscopic examination shall be fixed in 10 percent buffered formalin or a recognized suitable fixative as soon as necropsy is performed and no less than 48 hours prior to trimming.

16) Data and Reporting: Data shall be summarized in tabular form, showing for each test group the number of animals at the start of the test, the number of animals showing lesions, the types of lesions and the percentage of animals displaying each type of lesion. In addition to reporting requirements, the following specific information shall be reported:

- a. Test substance characterization shall include:
 - i. Chemical identification.
 - ii. Lot or batch number.
 - iii. Physical properties.
 - iv. Purity/impurities.
 - v. Identification and composition of any vehicle used.

- b. Test system shall contain data on:
 - i. Species and strain of animals used.
 - ii. Age including body weight data and sex.
 - iii. Test environment including cage conditions, ambient temperature, humidity, and light/dark periods.
 - iv. Identification of animal diet.
 - v. Acclimation period.
- c. Test procedure shall include the following data:
 - i. Method of randomization used.
 - ii. Full description of experimental design and procedure.
 - iii. Dose regimen including levels, methods, and volume.
- d. Test results shall include:
 - i. Group animal data. Tabulation of toxic response data by sex and exposure level for:
 - 1. Number of animals exposed.
 - 2. Number of animals showing signs of toxicity.
 - 3. Number of animals dying.
 - ii. Individual animal data. Data shall be presented as summary (group mean) as well as for individual animals.
 - 1. Date of death during the study or whether animals survived to termination.
 - 2. Date of observation of each abnormal sign and its subsequent course.
 - 3. Body weight data.
 - 4. Feed and water (if collected) consumption data.
 - 5. Dose (mg/kg/day).
 - 6. Results of toxicokinetic evaluation.
 - 7. Results of ophthalmological examination.
 - 8. Results of hematological tests performed.
 - 9. Results of clinical chemistry tests performed.
 - 10. Results of urinalysis, if performed.
 - 11. Necropsy findings, including absolute and relative (to body weight) organ weight data.
 - 12. Detailed description of all histopathological findings.
 - 13. Statistical treatment of results, where appropriate.

B. Deliverables

Within 11 weeks following necropsy in the 14-day repeated oral dose range finding study (Task 4.2), the contractor shall provide a report of the study results electronically to the EPA in MS Word format and all data in Excel spreadsheets.

1.7 TASK 5: REPEATED DOSE 90-DAY ORAL TOXICITY STUDY

Purpose: The purpose of Task 5 is for the contractor to conduct a GLP-compliant, repeated dose 90-day oral toxicity study in the rat. The contractor shall follow the EPA Health Effects Test Guideline OPPTS 870.3100 “90-Day Oral Toxicity in Rodents” August 1998 edition EPA 712-C-98-199 with modifications listed below in Section 5.1A. Studies may be conducted on up to six compounds. Formal study protocols are required.

Task 5 will be Firm Fixed Price.

Task 5.1: Repeated Dose 90-Day Oral Toxicity Study

A. Repeated Dose 90-Day Oral Toxicity Study

Modifications to the study design:

- 1) Species and Sex: Testing shall be performed in male and female Sprague-Dawley IGS rats from Charles Rivers Laboratory.
- 2) Age: Testing shall be performed with clinically healthy animals following a 7 – 10-day acclimation period. Dosing shall begin with animals no later than 8 – 9 weeks of age.
- 3) Control and Test Substance: The vehicle and test substances will be defined in Task 3.
- 4) Dose Levels: A total of 5 dose levels shall be used. Dose levels will be provided by USEPA based on the 14-day oral dose range finding study. The highest dose level shall result in toxic effects but not produce an incidence of fatalities which would prevent a meaningful evaluation. The intermediate dose levels shall be spaced to produce a gradation of toxic effects. The lowest dose level shall produce no evidence of toxicity. A concurrent vehicle control group is required.
- 5) Administration of Test Substance: The test substance shall be administered via oral gavage every day for 90 consecutive days. A dose volume of 5 ml/kg is preferred but 10 mL/kg is acceptable if required for solubility. The dose shall be given at approximately the same time each day and adjusted based on weekly body weights to maintain a consistent dose level.
- 6) No satellite groups shall be run in this study.
- 7) Plasma collection: At the time of sacrifice, blood shall be collected (using K3 EDTA as anticoagulant) by cardiac puncture under 70% CO₂:30% O₂ anesthesia. Plasma shall be isolated, stored at -80° C (6 x 100 µL aliquots of each). The time of blood collection and necropsy shall be recorded for all animals. Plasma samples analyzed by contractor for parent compound and metabolites by a suitable chromatographic/mass spectrometry method. The government's primary objective of this analysis is the quantitation of parent compound in plasma. Based on the method for the parent, we would like to identify, but not quantify, any metabolites that are present.
- 8) Sacrifice: At termination, all survivors in the treatment and control groups shall be weighed and then sacrificed. The order of sacrifice among animals shall be randomized.
- 9) Gross Necropsy: All animals shall be subjected to a full gross necropsy which includes examination of the external surface of the body, all orifices, and the cranial, thoracic and abdominal cavities and their contents.

- a. The liver, kidneys, lungs, adrenals, testes, epididymides, ovaries, uterus, thymus, spleen, brain, and heart shall be trimmed and weighed wet, as soon as possible after dissection.

B. Deliverables

Within 4 months of the necropsy in the subchronic study, the contractor shall provide a report of the study results electronically to the EPA in MS Word format and all data in Excel spreadsheets.

Task 5.2: Collect, Compile, Review Data and Prepare, Submit Reports

When all tasks for each test chemical have been completed or at the end of the POP, the contractor shall compile the data into one (1) electronic document. The written report, in MS Word, shall include analytical chemistry results of the stock, methods and results of the toxicity assays and analytical chemistry methods, and a section detailing any deviations or observed anomalies. The report must describe the methods and findings and shall include a comparison of test data to any control data with a description of significant findings.

QC shall be conducted on actual calculated numbers and not solely on data recording sheets. If problems are found in 10% of data that initially undergo a random check in accordance with the QC process, the remaining 90% of data shall be reviewed.

The data shall be provided to the TO COR electronically in an MS Excel spreadsheet or other similar electronic format. The data sheets shall be delivered to the TO COR after review by a/the senior scientist for data soundness and scientific relevance. All data shall be submitted (suspected outliers shall not be omitted from the report to EPA). The contractor shall ensure the overall quality of the final reports.

The draft report and data shall be delivered to the TO COR after review by the contractor senior scientist and the contractor Quality Assurance Manager (QAM) for scientific relevance and data soundness, respectively. These deliverables shall be as complete as possible and free of typographical errors and shall be provided as a cohesive unit that encompasses all assays performed for a specific chemical.

1.8 TASK 6 (OPTIONAL): REPEATED DOSE 28-DAY ORAL IMMUNOTOXICITY STUDY IN RATS

Purpose: The purpose of Task 6 is for the contractor to conduct a GLP-compliant repeated dose 28-day oral immunotoxicity study in rats. Contractor work on this task is dependent on the availability of funds. The contractor shall use the technical guidelines as described in the EPA Health Effects Guideline OPPTS 870.7800 Immunotoxicity June 1996, EPA 712-C-96-351, with modifications as described in Section 6.3(A). Formal study protocols are required.

Task 6 will be Firm-Fixed Price.

Task 6.1: Provide Revised Quality Assurance Project Plan and Standard Operating Procedures for 28-day Oral Immunotoxicity Study

The Contractor shall implement a quality system that meets ANSI standard E4-2014.

For planning purposes, assume that a Quality Assurance Project Plan (QAPP) will be required for Task 6). The contractor shall create a revised Quality Assurance Project Plan (QAPP) that documents

the planning, implementation, and assessment procedures for quality assurance and quality control activities. The QAPP integrates all the technical and quality aspects of the project to provide a blueprint for obtaining the type and quality of environmental data and information needed for a specific decision or use. The QAPP shall be prepared in accordance with the specifications identified by EPA (found at <https://www.epa.gov/quality/epa-qar-5-epa-requirements-quality-assurance-project-plans>).

- Within 10 business days after technical direction to begin optional Task 6, the contractor shall revise the QAPP that was previously developed for this task order and submit the revised QAPP to the TO COR for review.
- EPA will review the contractor's draft revised QAPP and provide the Contractor with written approval or written comments.
- If needed, the Contractor shall submit a revised QAPP within 5 business days of receipt of the written comments on the draft QAPP, unless otherwise instructed by the EPA TO COR.
- Under no circumstances shall work that involves the generation, collection, evaluation, analysis, or use of environmental data be performed by the contractor until the contractor receives written notification from the EPA TO COR that EPA has approved the contractor's QAPP.
- All QA documentation, including the QAPP, prepared under this Task Order, shall be considered non-proprietary, and shall be made available to the public upon request.
- The contractor also shall provide EPA with monthly reports of QA-related activities performed during implementation of this Task Order. These monthly QA reports shall identify QA activities performed to support implementation of this task order, problems encountered, deviations from the QAPP, and corrective actions taken.
- Within 2 months of acceptance of QAPP, provide draft SOPs for Task 6.2 and 6.3. EPA review, 15 business days; Vendor revision, 15 business days; sponsor approval, 15 business days.

Task 6.2: Determine optimum dose of sheep red blood cells administered and time to collect to collect sera.

A. Immunization with sheep red blood cells

- 1) Sheep red blood cells are used to immunize the rats in this study. Before conducting the main study, the optimum dose of sheep red blood cells and the time to collect blood from the immunized rats needs to be determined. Male and female Sprague-Dawley IGS rats from Charles Rivers Laboratory, 3 animals/sex/dose/time point, aged 10-12 weeks shall be used in this study. Three dose levels of SRBC's and three time points shall be used and determined after consultation between the contractor and EPA. The animals must be immunized by intravenous injection of sheep red blood cells. Control animals shall be treated with vehicle. The animals shall be monitored once per day for signs of any side effects from administration of the red blood cells. Moribund animals shall be euthanized. The animals shall be

euthanized on either 4, 5 or 6 days after administration of SRBC. Blood and spleen shall be collected. Spleen cells shall be isolated or serum shall be prepared and, either the plaque forming cell (PFC) assay (measures IgM response) or an enzyme linked immunosorbent assay (ELISA) (measures IgM levels) must be performed, respectively, to determine the optimum concentration of SRBCs and sacrifice time for the immunotoxicology study. See Temple et al. (1993, 1995) for more details in References below. Decision on the assay shall be made in consultation with EPA.

2) Report of results shall include:

- a. Test substance characterization
 - i. Source of sheep red blood cells and lot number if available.
- b. Test system
 - i. Source of animals with age, weight and sex
 - ii. Animal husbandry conditions
 - iii. Diet
- c. Experimental procedure
 - i. Method of randomization of animals
 - ii. Full description of experimental design and procedure
 - iii. Dose regimen
- d. Test results
 - i. Group animal data
 - 1. Number of animals exposed
 - 2. Number of animals displaying adverse effect
 - a) Description of the adverse effects
 - 3. Number of animals dying
 - 4. Results of IgM response or concentration, depending on the test conducted.
 - 5. The report shall include concentration of SRBC for either the plaque forming cell (PFC) assay (measures IgM response) or an enzyme linked immunosorbent assay (ELISA) (measures IgM levels) at three time points

B. Deliverable

Within one month of completion of study, the contractor shall provide a report of the study results electronically to the EPA in MS Word format and all data in Excel spreadsheets.

Task 6.3: Repeated Dose 28-day Oral Immunotoxicity Study.

A. Oral Dose Immunotoxicity Study

Modifications to the study design:

- 1) Species and Sex: Testing shall be performed in Sprague-Dawley IGS rats from Charles Rivers Laboratory. The sex of the animals used in the study will be selected by EPA.
- 2) Control and Test Substance: The vehicle and test substances will be defined in Task 3.
- 3) Dose Levels: A total of 5 dose levels shall be used. Dose levels will be provided by USEPA based either on the 14-day oral dose range finding study or the 90-day oral toxicity study.

The highest dose shall not produce significant stress, malnutrition or fatalities, but produce some sign of measurable general toxicity (e.g., ten percent loss of body weight). A concurrent vehicle control group and positive control (cyclophosphamide) are required.

- 4) Administration of Test Substance: The test substance shall be administered via oral every day for 28 consecutive days. A dose volume of 5 ml/kg is preferred but 10 mL/kg is acceptable if required for solubility. The dose shall be given at approximately the same time each day and adjusted based on weekly body weights to maintain a consistent dose level.
- 5) Rats shall be immunized with SRBC's based on the findings from Task 6.2 and CCTE recommendations.
- 6) Blood and spleen shall be collected. Spleen cells shall be isolated or serum shall be prepared and, either the plaque forming cell (PFC) assay (measures IgM response) or an enzyme linked immunosorbent assay (ELISA) (measures IgM levels) must be performed, respectively, to determine the optimum concentration of SRBCs and sacrifice time for the immunotoxicology study. See Temple et al. (1993, 1995) for more details in References below. Decision on the assay shall be made in consultation with EPA.

B. Deliverables

Within 4 months of necropsy in the immunotoxicity study, the contractor shall provide a report of the study results electronically to the EPA in MS Word format and all data in Excel spreadsheets.

1.9 REPORTING REQUIREMENTS AND DELIVERABLES

The contractor shall provide the following deliverables listed in Table 2.

Table 1: Deliverables and Schedule

Tasks	Deliverables	Due Dates
Task 1	Task management <ul style="list-style-type: none"> ✓ Kick-off meeting within 2 weeks after the task order has been awarded. ✓ Monthly Progress Report (i.e. Technical/Progress Status Report and Financial Status Report) by the 15th of each month (following completion of 1st reporting period). ✓ Biweekly status meetings ✓ Other meetings as required by the TOCOR ✓ Email a copy of Monthly Progress Report to the CO, Contract level COR, TOCOR, and Alt TOCOR ✓ Immediately inform the TOCOR when any hours or costs for any task has exceeded or is expected to exceed the contractor estimate by >10%. ✓ Immediately inform the TOCOR of any problems that may impact the production, budget, and/or delivery of deliverables. ✓ The Contractor shall notify the TOCOR when 75% of the Government approved hours or approved LH costs have been incurred (including unbilled hours and costs). 	
Task 2	EPA Requirements for Quality Assurance Project Plans (QA/R- 5) https://www.epa.gov/quality/epa-qar-5-epa-requirements-quality-assurance-project-plans	<ul style="list-style-type: none"> ✓ Draft QAPP. Within 10 business days after Task Order Award. ✓ Final QAPP. Within 5 business days of receipt of the written

Tasks	Deliverables	Due Dates
	<ul style="list-style-type: none"> ✓ Draft QAPP ✓ Final QAPP 	comments on the draft QAPP, unless otherwise instructed by the EPA TO COR.
Task 3.1	<ul style="list-style-type: none"> ✓ Provide EPA information regarding test chemical procured. Include purity, lot number, and other information provided by the manufacturer. Information provided electronically in Word format. 	<ul style="list-style-type: none"> ✓ Within two weeks of acquisition of chemical.
Task 3.3	<ul style="list-style-type: none"> ✓ Draft SOPs shall be provided to the EPA/EPA for approval. Each SOP shall have clearly labeled sections similar to: 1.0 Purpose; 2.0 Scope; 3.0 Responsibilities; 4.0 Procedures; 5.0 References. The draft SOPs shall be delivered to the EPA electronically in MS Word format and all data shall be provided in Excel spreadsheets. ✓ The analytical chemistry methods must be finalized. For each test chemical, a brief report shall be provided to the EPA which outlines the method and clearly indicates the Limit of Quantitation (LOQ) and Level of Detection (LOD) achieved with that method. Purity of test chemicals shall be reported and must >95%. The levels of the potential contaminants perfluorooctanoic acid (PFOA) and perfluorooctane sulfonic acid shall be reported. This report shall be in a Word format with all data provided in an Excel format. 	<ul style="list-style-type: none"> ✓ Draft SOP within two (2) months approval of QAPP ✓ Chemical purity determination within 8 weeks after acceptance of chemistry SOPs.
Task 3.3	<ul style="list-style-type: none"> ✓ Report on solubility and stability of test chemical in vehicle. Report shall include vehicles tested, methods used to measure solubility and stability, solubility, stability of the test chemical in the vehicle for 7 days. This report shall be in a Word format with all data provided in an Excel format. 	<ul style="list-style-type: none"> ✓ Within three (3) months of acceptance of QAPP.
Task 4.1	<ul style="list-style-type: none"> ✓ Report on oral tolerability study. Report shall be in MS Word format and data in Excel spreadsheet. 	<ul style="list-style-type: none"> ✓ Within two (2) weeks of necropsy.
Task 4.2	<ul style="list-style-type: none"> ✓ Report on 14-day repeated dose oral toxicity study. Report shall be in MS Word format and data in Excel spreadsheet. 	<ul style="list-style-type: none"> ✓ Within eleven (11) weeks of completion of necropsy.

Tasks	Deliverables	Due Dates
Task 5.1	✓ Report on 90-day repeated dose oral toxicity study. Report shall be in MS Word format and data in Excel spreadsheet.	✓ Within four (4) months of necropsy.
Task 6.1	✓ Draft QAPP	✓ Within 10 business days after Technical Direction.
Task 6.2	✓ Report on concentration of SRBCs to use in immunotoxicology study	✓ Within one (1) month of completion of study
Task 6.3	✓ Report of the study results electronically to the EPA in MS Word format and all data in Excel spreadsheets.	✓ Within 4 months of necropsy in the immunotoxicity study
Task 3,4,5,6	✓ Any unused test substance, residual wet tissues, pathology blocks, slides or residual tissues shall be shipped to the USEPA. The SRBCs can be disposed of by the contractor.	✓ After acceptance of final reports for each task

1.10 ACCEPTABLE QUALITY LEVEL FOR TASKS

See Attachment 2: Quality Assurance Surveillance Plan

1.11 PERIOD OF PERFORMANCE

The period of performance of this task order is:

BASE: 36 months from award date.

1.12 PLACE OF PERFORMANCE

Work shall be performed off-site.

1.13 PERSONNEL

The Contractor is responsible for providing personnel with the necessary level expertise to support the task activities and requirement in this PWS.

1.14 TASK ORDER TYPE: TIME & MATERIALS AND FIRM FIXED PRICE

1.15 GOVERNMENT FURNISHED PROPERTY (GFP)

In accordance with FAR 45.102, the contractor shall furnish all property required for performing Government contracts. If a contractor believes that Government property is required for performance of the contract, the contractor shall submit a written request to the CO. For cost purposes, assume that EPA shall provide an office phone with voicemail, and e-mail for approved personnel working in OSCP-space to complete work under Task 3 of this task order.

1.16 TRAVEL

The Contractor may be required to travel in the course of the performance of this task order. The Contractor is required to follow the requirements of subpart 31.2 of the FAR regulations in incurring allowable travel costs under this task order, and correspondingly must, at all times, seek and obtain government rates whenever available and observe current subsistence ceilings.

1.17 TRAINING

EPA-H-31-105 APPROVAL OF TRAINING [see Section H.22 of the ID/IQ contract]

- (a) The contractor shall provide and maintain a qualified staff of personnel to meet the requirements of the Performance Work Statement. The contractor shall provide training to keep its personnel abreast of changes to the science and/or technology associated with the requirements of the contract. In addition, the contractor shall ensure that its personnel receive appropriate safety, health and environmental training in accordance with Federal, state and local requirements prior to assigning any task that require such training. The contractor shall provide documentation of such training upon the request of the Contract-Level COR and/or Contracting Officer.

The Government will not directly reimburse the cost for contractor employees to meet or maintain minimal contract requirements or to obtain and sustain an appropriate level of professionalism. Any direct charges for training will only be considered for reimbursement under this contract by compliance with the procedures set forth in paragraph (b) (see Section H.22 of the IDIQ contract).

2 INSPECTION AND ACCEPTANCE

QUALITY ASSURANCE PROJECT PLAN (SEE TASK 2 ABOVE)

The Contractor shall submit a draft QAPP per EPA Requirements for Quality Assurance Project Plans (QA/R-5)

Table 3: Quality Assurance Project Plan

	Documentation	Specifications	Due
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X	Quality Assurance Project Plan for the Task Order	EPA Requirements for Quality Assurance Project Plans (QA/R-5) (dated 3//20/2011) https://www.epa.gov/quality/epa-qar-5-epa-requirements-quality-assurance-project-plans	<ul style="list-style-type: none"> ✓ Draft QAPP. Within 10 business days after Task Order Award. ✓ Final QAPP. Within 5 business days of receipt of the written comments on the draft QAPP, unless otherwise instructed by the EPA TO COR.
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3 TASK ORDER ADMINISTRATION DATA

3.1 CONTRACT ADMINISTRATION REPRESENTATIVES (EPA LOCAL CLAUSE EPA-G-42-101)

- Administrative Contracting Officer (ACO):
Sean Gifford
U.S. EPA
26 West Martin Luther King Drive
Cincinnati, OH 45268
Phone: 513-487-2506
Email: Gifford.Sean@epa.gov

- Contract Level Contracting Officer's Representative (CLCOR):
Cathy Stewart
U.S. EPA
2777 S Crystal Drive
Mail Code: 7502P
Arlington, VA 22202
Phone: 703-305-7711
Email: Stewart.Cathy@epa.gov

- Alternate CLCOR:
LaTangila Edwards
U.S. EPA
2777 S Crystal Drive
Mail Code: 7509P
Arlington, VA 22202
Phone: 703-305-7170
Email: Edwards.Latangila@epa.gov

- Task Order Contracting Officer's Representative (TOCOR):
Patrice Borsz
U.S. EPA
109 T.W. Alexander Drive
Mail Code: E205-09
Research Triangle Park, NC 27711
Phone: 919-541-5233
Email: Borsz.Patrice@epa.gov

- Alternate TOCOR:
Jenna Tomayko
U.S. EPA
109 T.W. Alexander Drive
Mail Code: E205-09
Research Triangle Park, NC 27711
Phone: 919-541-2538
Email: Tomayko.Jenna@epa.gov

3.2 TASK ORDER CLAUSES

EPAAR 1552.237-72 KEY PERSONNEL (APR 1984)

- (a) The Contractor shall assign to this contract the following key personnel:

Lead Chemist: (b) (4)

Lead Pathologist: (b) (4)

Study Director: (b) (4)

- (b) During the first ninety (90) days of performance, the Contractor shall make no substitutions of key personnel unless the substitution is necessitated by illness, death, or termination of employment. The Contractor shall notify the Contracting Officer within 15 calendar days after the occurrence of any of these events and provide the information required by paragraph (c) of this clause. After the initial 90-day period, the Contractor shall submit the information required by paragraph (c) to the Contracting Officer at least 15 days prior to making any permanent substitutions.
- (c) The Contractor shall provide a detailed explanation of the circumstances necessitating the proposed substitutions, complete resumes for the proposed substitutes, and any additional information requested by the Contracting Officer. Proposed substitutes should have comparable qualifications to those of the persons being replaced. The Contracting Officer will notify the Contractor within 15 calendar days after receipt of all required information of the decision on substitutions. This clause will be modified to reflect any approved changes of key personnel.
(End of clause)

References

- Cornacoff, J.B., Graham, C.S., and LaBrie, T.K. 1995. Phenotypic identification of peripheral blood mononuclear leukocytes by flow cytometry as an adjunct to immunotoxicity evaluation. In *Methods in Immunotoxicology* (G.R. Burleson, J.H. Dean, and A.E. Munson, Eds.), Vol. 1, pp 211–226, Wiley-Liss, Inc., New York.
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- U.S. EPA Health Effects Test Guideline OPPTS 870.3100 90-Day Oral Toxicity in Rodents
- U.S. EPA Health Effects Test Guideline OPPTS 870.7800 Immunotoxicity

ATTACHMENT 2

QUALITY ASSURANCE SURVEILLANCE PLAN FOR TASK ORDERS

QUALITY ASSURANCE SURVEILLANCE PLAN

PERFORMANCE REQUIREMENT	PERFORMANCE MEASURE (PM)	PERFORMANCE STANDARD	SURVEILLANCE METHOD	INCENTIVES & DISINCENTIVES
<p><u>MANAGEMENT AND COMMUNICATION:</u></p> <p>The contractor shall maintain contact with the EPA Contracting Officer (CO), Contracting Officer Representative (COR), and Task Order Contracting Officer Representative (TOCOR) throughout the performance of the contract.</p>	<p>Contractor shall immediately bring potential problems to the appropriate EPA personnel and shall recommend actions that would mitigate or resolve the problem.</p>	<p>Issues that impact project schedules and costs shall be brought to the attention of the EPA within 3- business days of occurrence.</p>	<p>All active task orders will be reviewed by the EPA to identify unreported issues.</p>	<p>Performance will be considered in the award of subsequent task orders and will be factored into the annual evaluation of Business Relations in the Contractor Performance Assessment Reporting System (CPARS).</p>
<p><u>TIMELINESS:</u></p> <p>For every Task Order awarded establishing a firm, specific delivery date for the generation of a report, the contractor shall deliver such report to the COR, TOCOR and CO no later than the time specified in the order's PWS.</p>	<p>Deliverables and related work must comply with contractual timeliness requirements. The contractor will be evaluated on its responsiveness to all task orders.</p>	<p>95% of all deliverables and related work shall be completed on time within task schedule and/or tech. direction requirements.</p>	<p>100% inspection of all deliverables and related work by the TOCOR; TOCOR will document the timeliness of all work requirements.</p>	<p>Performance will be considered in the award of subsequent task orders and will be factored into the annual evaluation of Timeliness in the Contractor Performance Assessment Reporting System (CPARS).</p>
<p><u>TECHNICAL QUALITY:</u></p> <p>For every task order awarded, the analyses conducted by the contractor shall be factual, defensible, credible, and based on sound scientific methods. All data shall be collected from reputable sources and quality assurance measures shall be conducted in accordance with the agency requirements outlined in the task orders.</p>	<p>All deliverables and related work must be complete, accurate, thorough, and professionally credible.</p>	<p>Data are 100% accurate; review demonstrates a high level of expertise and credibility with regard to personnel and use of scientific methodology. Task Orders shall be conducted in strict conformance with approved QA plans. Outputs shall withstand internal review by the US EPA and outside scientific reviewers.</p>	<p>EPA Staff will conduct secondary reviews of work completed by the contractor. Feedback will be provided.</p>	<p>Performance will be considered in the award of subsequent task orders and will be factored into the annual evaluation in the category of Quality of Product or Service in the Contractor Performance Assessment Reporting System (CPARS).</p>

NAME OF OFFEROR OR CONTRACTOR
 Research Triangle Institute

ITEM NO. (A)	SUPPLIES/SERVICES (B)	QUANTITY (C)	UNIT (D)	UNIT PRICE (E)	AMOUNT (F)
	LIST OF CHANGES: Reason for Modification: Other Administrative Action Payment: RTP Finance Center US Environmental Protection Agency RTP-Finance Center (AA216-01) 109 TW Alexander Drive www2.epa.gov/financial/contracts Durham NC 27711 Period of Performance: 06/09/2021 to 06/08/2024				

AMENDMENT OF SOLICITATION/MODIFICATION OF CONTRACT				1. CONTRACT ID CODE		PAGE OF PAGES	
						1 2	
2. AMENDMENT/MODIFICATION NO.		3. EFFECTIVE DATE		4. REQUISITION/PURCHASE REQ. NO.		5. PROJECT NO. (If applicable)	
P00002		See Block 16C		PR-ORD-21-02528			
6. ISSUED BY		CODE		7. ADMINISTERED BY (If other than Item 6)		CODE	
CAD							
US Environmental Protection Agency 26 West Martin Luther King Drive Mail Code: W136 Cincinnati OH 45268-0001							
8. NAME AND ADDRESS OF CONTRACTOR (No., street, county, State and ZIP Code)				(x)			
Research Triangle Institute Attn: Linda Eubanks PO BOX 12194 RESEARCH TRIANGLE PARK NC 277092194							
				9A. AMENDMENT OF SOLICITATION NO.			
				9B. DATED (SEE ITEM 11)			
				x			
				10A. MODIFICATION OF CONTRACT/ORDER NO.			
				68HE0H18D0008			
				68HERC21F0259			
				10B. DATED (SEE ITEM 13)			
				06/08/2021			
CODE 004868105		FACILITY CODE					
11. THIS ITEM ONLY APPLIES TO AMENDMENTS OF SOLICITATIONS							
<input type="checkbox"/> The above numbered solicitation is amended as set forth in Item 14. The hour and date specified for receipt of Offers <input type="checkbox"/> is extended. <input type="checkbox"/> is not extended. Offers must acknowledge receipt of this amendment prior to the hour and date specified in the solicitation or as amended, by one of the following methods: (a) By completing Items 8 and 15, and returning _____ copies of the amendment; (b) By acknowledging receipt of this amendment on each copy of the offer submitted; or (c) By separate letter or electronic communication which includes a reference to the solicitation and amendment numbers. FAILURE OF YOUR ACKNOWLEDGEMENT TO BE RECEIVED AT THE PLACE DESIGNATED FOR THE RECEIPT OF OFFERS PRIOR TO THE HOUR AND DATE SPECIFIED MAY RESULT IN REJECTION OF YOUR OFFER. If by virtue of this amendment you desire to change an offer already submitted, such change may be made by letter or electronic communication, provided each letter or electronic communication makes reference to the solicitation and this amendment, and is received prior to the opening hour and date specified.							
12. ACCOUNTING AND APPROPRIATION DATA (If required)							
See Schedule							
13. THIS ITEM ONLY APPLIES TO MODIFICATION OF CONTRACTS/ORDERS. IT MODIFIES THE CONTRACT/ORDER NO. AS DESCRIBED IN ITEM 14.							
CHECK ONE	A. THIS CHANGE ORDER IS ISSUED PURSUANT TO: (Specify authority) THE CHANGES SET FORTH IN ITEM 14 ARE MADE IN THE CONTRACT ORDER NO. IN ITEM 10A.						
X	B. THE ABOVE NUMBERED CONTRACT/ORDER IS MODIFIED TO REFLECT THE ADMINISTRATIVE CHANGES (such as changes in paying office, appropriation data, etc.) SET FORTH IN ITEM 14, PURSUANT TO THE AUTHORITY OF FAR 43.103(b).						
	C. THIS SUPPLEMENTAL AGREEMENT IS ENTERED INTO PURSUANT TO AUTHORITY OF:						
	D. OTHER (Specify type of modification and authority)						
E. IMPORTANT: Contractor <input checked="" type="checkbox"/> is not <input type="checkbox"/> is required to sign this document and return _____ copies to the issuing office.							
14. DESCRIPTION OF AMENDMENT/MODIFICATION (Organized by UCF section headings, including solicitation/contract subject matter where feasible.)							
DUNS Number: 004868105+7709 Endocrine Disruptor Screening Program Technical and Support Services ID/IQ Task Order: Laboratory Support for Repeated Dose Oral Toxicity Studies of Poly- and Perfluoroalkyl Substances in Rats TOCOR: Linda Adams Max Expire Date: 06/08/2024 Invoice Approver: Linda Adams Alt Invoice App: Jenna Tomayko This unilateral Modification P00002 is issued to change the primary TOCOR from Patrice Borsz to Linda Adams. The new TOCOR's contact information is provided below. All other contract administration representatives remain unchanged.							
Linda Adams Continued ...							
Except as provided herein, all terms and conditions of the document referenced in Item 9 A or 10A, as heretofore changed, remains unchanged and in full force and effect.							
15A. NAME AND TITLE OF SIGNER (Type or print)				16A. NAME AND TITLE OF CONTRACTING OFFICER (Type or print)			
				Sean Gifford			
15B. CONTRACTOR/OFFEROR		15C. DATE SIGNED		16B. UNITED STATES OF AMERICA		16C. DATE SIGNED	
				 (Signature of Contracting Officer)		ELECTRONIC SIGNATURE 09/28/2021	
(Signature of person authorized to sign)							

CONTINUATION SHEET	REFERENCE NO. OF DOCUMENT BEING CONTINUED 68HE0H18D0008/68HERC21F0259/P00002	PAGE	OF
		2	2

NAME OF OFFEROR OR CONTRACTOR
Research Triangle Institute

ITEM NO. (A)	SUPPLIES/SERVICES (B)	QUANTITY (C)	UNIT (D)	UNIT PRICE (E)	AMOUNT (F)
	<p>U.S. EPA 109 T.W. Alexander Drive Mail Code B105-03 Research Triangle Park, NC 27711 Phone: 919-541-1424 Email: Adams.Lindad@epa.gov</p> <p>All other terms and conditions remain unchanged and in full force and effect.</p> <p>LIST OF CHANGES: Reason for Modification: Other Administrative Action Task Order Contracting Officer Representative changed from Patrice Borsz to Linda Adams Payment: RTP Finance Center US Environmental Protection Agency RTP-Finance Center (AA216-01) 109 TW Alexander Drive www2.epa.gov/financial/contracts Durham NC 27711 Period of Performance: 06/09/2021 to 06/08/2024</p>				

AMENDMENT OF SOLICITATION/MODIFICATION OF CONTRACT		1. CONTRACT ID CODE		PAGE OF PAGES 1 3	
2. AMENDMENT/MODIFICATION NO. P00003		3. EFFECTIVE DATE See Block 16C		5. PROJECT NO. (If applicable)	
6. ISSUED BY CAD US Environmental Protection Agency 26 West Martin Luther King Drive Mail Code: W136 Cincinnati OH 45268-0001		7. ADMINISTERED BY (If other than Item 6) CODE		9A. AMENDMENT OF SOLICITATION NO.	
8. NAME AND ADDRESS OF CONTRACTOR (No., street, county, State and ZIP Code) Research Triangle Institute Attn: Linda Eubanks PO BOX 12194 RESEARCH TRIANGLE PARK NC 277092194		(x)		9B. DATED (SEE ITEM 11)	
CODE JJHCMK4NT5N3		FACILITY CODE		10A. MODIFICATION OF CONTRACT/ORDER NO. 68HE0H18D0008 68HERC21F0259 10B. DATED (SEE ITEM 13) 06/08/2021	
11. THIS ITEM ONLY APPLIES TO AMENDMENTS OF SOLICITATIONS					
<input type="checkbox"/> The above numbered solicitation is amended as set forth in Item 14. The hour and date specified for receipt of Offers <input type="checkbox"/> is extended. <input type="checkbox"/> is not extended. Offers must acknowledge receipt of this amendment prior to the hour and date specified in the solicitation or as amended, by one of the following methods: (a) By completing Items 8 and 15, and returning _____ copies of the amendment; (b) By acknowledging receipt of this amendment on each copy of the offer submitted; or (c) By separate letter or electronic communication which includes a reference to the solicitation and amendment numbers. FAILURE OF YOUR ACKNOWLEDGEMENT TO BE RECEIVED AT THE PLACE DESIGNATED FOR THE RECEIPT OF OFFERS PRIOR TO THE HOUR AND DATE SPECIFIED MAY RESULT IN REJECTION OF YOUR OFFER. If by virtue of this amendment you desire to change an offer already submitted, such change may be made by letter or electronic communication, provided each letter or electronic communication makes reference to the solicitation and this amendment, and is received prior to the opening hour and date specified.					
12. ACCOUNTING AND APPROPRIATION DATA (If required) See Schedule					
13. THIS ITEM ONLY APPLIES TO MODIFICATION OF CONTRACTS/ORDERS. IT MODIFIES THE CONTRACT/ORDER NO. AS DESCRIBED IN ITEM 14.					
CHECK ONE					
A. THIS CHANGE ORDER IS ISSUED PURSUANT TO: (Specify authority) THE CHANGES SET FORTH IN ITEM 14 ARE MADE IN THE CONTRACT ORDER NO. IN ITEM 10A.					
B. THE ABOVE NUMBERED CONTRACT/ORDER IS MODIFIED TO REFLECT THE ADMINISTRATIVE CHANGES (such as changes in paying office, appropriation data, etc.) SET FORTH IN ITEM 14, PURSUANT TO THE AUTHORITY OF FAR 43.103(b).					
C. THIS SUPPLEMENTAL AGREEMENT IS ENTERED INTO PURSUANT TO AUTHORITY OF: X Adding instructions to the contractor					
D. OTHER (Specify type of modification and authority)					
E. IMPORTANT: Contractor <input checked="" type="checkbox"/> is not <input type="checkbox"/> is required to sign this document and return _____ copies to the issuing office.					
14. DESCRIPTION OF AMENDMENT/MODIFICATION (Organized by UCF section headings, including solicitation/contract subject matter where feasible.) UEI: JJHCMK4NT5N3+7709 Endocrine Disruptor Screening Program Technical and Support Services ID/IQ Task Order: Laboratory Support for Repeated Dose Oral Toxicity Studies of Poly- and Perfluoroalkyl Substances in Rats TOCOR: Linda Adams Max Expire Date: 06/08/2024 Invoice Approver: Linda Adams Alt Invoice App: Jenna Tomayko The purpose of this modification is to add instructions to the contractor on Line Item #3 of this TO. Chemical Procurement; the TOCOR shall instruct Contractor to purchase additional chemicals to develop analytical chemistry methods under EPA 68HEOH18D0008 - Task Order 68HERC21F0259 TO-5 with RTI; The EPA TOCOR. Current chemical of interest to purchase is listed below: Continued ... Except as provided herein, all terms and conditions of the document referenced in Item 9 A or 10A, as heretofore changed, remains unchanged and in full force and effect.					
15A. NAME AND TITLE OF SIGNER (Type or print)			16A. NAME AND TITLE OF CONTRACTING OFFICER (Type or print)		
			Mark Cranley		
15B. CONTRACTOR/OFFEROR		15C. DATE SIGNED		16B. UNITED STATES OF AMERICA	
(Signature of person authorized to sign)				ELECTRONIC SIGNATURE	
				16C. DATE SIGNED	
				12/13/2022	

CONTINUATION SHEET	REFERENCE NO. OF DOCUMENT BEING CONTINUED	PAGE	OF
	68HEOH18D0008/68HERC21F0259/P00003	2	3

NAME OF OFFEROR OR CONTRACTOR
Research Triangle Institute

ITEM NO. (A)	SUPPLIES/SERVICES (B)	QUANTITY (C)	UNIT (D)	UNIT PRICE (E)	AMOUNT (F)
	<p>Product Cas Number Chemical name: 2121-7-02 6189-02-2 2-Chlorotetrafluoropropionic Acid. All other terms and conditions remain unchanged. LIST OF CHANGES: Reason for Modification: Change Order Total Amount for this Modification: \$0.00 New Total Amount for this Version: \$717,549.00 New Total Amount for this Award: \$4,527,202.00 BIB - Infrastructure Bill Funds changed to : N/A CHANGES FOR LINE ITEM NUMBER: 3 Description changed from Base Period (Optional Task 6) in accordance with Attachment 1, Performance Work Statement.</p> <p>This is a firm-fixed priced contract line item number (CLIN).</p> <p>This CLIN has an associated period of performance from June 9, 2021 to June 8, 2024.</p> <p>This optional CLIN is exercised upon award of the task order. to Base Period (Optional Task 6) in accordance with Attachment 1, Performance Work Statement.</p> <p>This is a firm-fixed priced contract line item number (CLIN).</p> <p>This CLIN has an associated period of performance from June 9, 2021 to June 8, 2024.</p> <p>This optional CLIN is exercised upon award of the task order. The purpose of this modification is to add this below mentioned clause to this subject Task Order. Chemical Procurement; the TOCOR shall instruct Contractor to purchase additional chemicals to develop analytical chemistry methods under EPA 68HEOH18D0008 - Task Order 68HERC21F0259 TO-5 with RTI; The EPA TOCOR. Current chemical of interest to purchase is listed below: Product Cas Number Chemical name: 2121-7-02 6189-02-2 2-Chlorotetrafluoropropionic Acid. All other terms and conditions remain unchanged.</p> <p>Continued ...</p>				

NAME OF OFFEROR OR CONTRACTOR
Research Triangle Institute

ITEM NO. (A)	SUPPLIES/SERVICES (B)	QUANTITY (C)	UNIT (D)	UNIT PRICE (E)	AMOUNT (F)
	Payment: RTP Finance Center US Environmental Protection Agency RTP-Finance Center (AA216-01) 109 TW Alexander Drive www2.epa.gov/financial/contracts Durham NC 27711 Period of Performance: 06/09/2021 to 06/08/2024				